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seco" service under 37 CFR 1.10 on the date	LLC DOX nt and Trade	roved for use through 04/30/2003. OMB 0651/00 mark Office; U.S. DEPARTMENT OF COMMER
d abovended the Raderson to Rhou Chromatis of 1986, 1986 persons are red	Application Number	09/541,848
For	Filing Date	April 3, 2000
Continued Examination (RCE) Transmittal Address to: Commissioner for Patents P. O. Box 1450, Mail Stop RCE Alexandria, VA 22313-1450	First Named Inventor	Chen
	Art Unit	1635
	Examiner Name	Schultz, James
	Attorney Docket Number	HYB-015US4 (1006/016)
This is a Request for Continued Examination (RCE Request for Continued Examination (RCE) practice under 37 1995, or to any design application. See Instruction Sheet for	CFR 1.114 does not apply to any uti	lity or plant application filed prior to June 8,
1. (Submission required under 37 CFR 1.114)		
a. Previously submitted ; Consider the amendment(s)/reply und	ler 37 CFR 1 116 previously filed on	May 2, 2003 (copy enclosed)
i. (Any unentered amendment(s) referred to a		<u> </u>
li. Consider the arguments in the Appeal	Brief or Rely Brief previously filed or	1
b. Enclosed		
I. Amendment/Reply	iii. Information	Disclosure Statement (IDS)
ii. Affidavit(s)/ Declaration(s)	iv. Other	
Miscellaneous Suspension of action on the above-identified period of months. (Period of suspension of the above-identified period of months.)	ension shall not exceed 3 months; Fee und	* *
3. Fees The RCE fee under 37 CFR 1.17(e) is required. The Director is hereby authorized to charge		is filed.
a. Deposit Account No. 50-2285	e the following fees, or credit any ove	rpayments, to
	(0)	rpayments, to HBIZUNES 00000063 09541848
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a. Deposit Account No. 50-2285 i. RCE fee required under 37 CFR 1.176 ii. Extension of time fee (37 CFR 1.136 and 11) Other	(e) 08/20/2003 d 1.17) 01 FC::2801 enclosed	HBIZUNES 00000063 09541848
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Page I of 2

This collection of information is required by 37 CFR 1.114. The information is required to obtain or retain a benefit by the public which is to file (and by the USPTO to process) an application. Confidentiality is governed by 35 U.S.C. 122 and 37 CFR 1.14. This collection is estimated to take 12 minutes to complete, including gathering, preparing, and submitting the completed application form to the USPTO. Time will vary depending upon the individual case. Any comments on the uspect of time you require to complete this form and/or suggestions for reducing this burden, should be sent to the Chief Information Officer, U.S. Patent and Trademark Office, U.S. Department of Commerce, Washington, DC 20231. DO NOT SEND FEES OR COMPLETED FORMS TO THIS ADDRESS. SEND TO:

Commissioner for Patents, Box RCE, Washington, DC 20231.

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I THE UNITED STATES PATENT AND TRADEMARK OFFICE

IN APPLICATION OF:

ART UNIT: 1635

CHEN et al.

EXAMINER: Wang, Andrew J.

SERIAL NO.: 09/541,848

FILED: April 3, 200

FOR: MDM2 SPECIFIC ANTISENES

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CERTIFICATE OF FIRST CLASS MAILING UNDER 37 C.F.R. § 1.8

I hereby certify that this correspondence is being deposited with the United States Postal Service with sufficient postage as First Class Mail in an envelope addressed to: Commissioner for Patents, Washington, DC 20231 on the date indicated below.

Date: 4/22/03

Signature McMAA Sumpton

Printed Name: Melissa A. Simpson

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Assistant Commissioner for Patents Washington, DC 20231

AMENDMENT AND RESPONSE UNDER 37 C.F.R. § 1.116

Dear Sir:

In response to the Final Office Action dated October 22, 2002, please reconsider the above-referenced patent application in view of the following amendment and remarks. Included with this response is a Request for an Extension of Time for 3 months under 37 C.F.R. § 1.136(a) and the appropriate fee.

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IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

<u>AMENDMENTS</u>

LISTING OF THE CLAIMS

- 1. (Presently Amended) A method of inhibiting expression of MDM2 in a mammal, the method comprising administering to the mammal an effective MDM2-expression inhibiting amount of an anti-MDM2 antisense oligonucleotide, wherein said antisense oligonucleotide comprises from about 8 to about 50 nucleotides that inhibits MDM2 protein expression, said oligonucleotide binding to mdm2-encoding RNA and being complementary to a sequence that overlaps by at least one nucleotide a sequence within the mdm2 RNA, which sequence within the mdm2 RNA is selected from the group consisting of SEQ ID NOS: 2, 3, 4, 7, 8, 9, 10, 11, 13, 14, 15, 16, 17, 18, 19, 20, 21, 22, 23, and 24.
- 2. The method according to claim 1 comprising co-administering a cancer chemotherapeutic agent.
- 3. The method according to claim 2, wherein the cancer therapeutic agent is 10-hydorxycamptothecin, adriamycin, or 5-fluorouracil.
- 4. The method according to claim 1 comprising co-treating the mammal with anti-cancer levels of radiation.
- 5. (Presently Amended) A method of inhibiting cancer *in vivo*, the method comprising administering a cancer-inhibiting amount of an anti-MDM2 antisense oligonucleotide, wherein the cancer involves over expression of MDM2, wherein said antisense oligonucleotide comprises from about 8 to about 50 nucleotides that inhibits MDM2 protein expression, said oligonucleotide binding to mdm2-encoding RNA and being complementary to a sequence that overlaps by at least one nucleotide a sequence within the mdm2 RNA, which sequence within the mdm2 RNA is selected from the